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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/874,991	06/07/2001	James J. Mond	07787.0042	5537

22852 7590 09/08/2004

FINNEGAN, HENDERSON, FARABOW, GARRETT & DUNNER
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1300 I STREET, NW
WASHINGTON, DC 20005

EXAMINER

MINNIFIELD, NITA M

ART UNIT	PAPER NUMBER
	1645

DATE MAILED: 09/08/2004

Please find below and/or attached an Office communication concerning this application or proceeding.

Office Action Summary	Application No.	Applicant(s)	
	09/874,991	MOND ET AL.	
	Examiner N. M. Minnifield	Art Unit 1645	

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --
Period for Reply

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) FROM
THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

Status

1) Responsive to communication(s) filed on 10 June 2004.
 2a) This action is FINAL. 2b) This action is non-final.
 3) Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

Disposition of Claims

4) Claim(s) 1-17 is/are pending in the application.
 4a) Of the above claim(s) 11-17 is/are withdrawn from consideration.
 5) Claim(s) _____ is/are allowed.
 6) Claim(s) 1-10 is/are rejected.
 7) Claim(s) _____ is/are objected to.
 8) Claim(s) 11-17 are subject to restriction and/or election requirement.

Application Papers

9) The specification is objected to by the Examiner.
 10) The drawing(s) filed on _____ is/are: a) accepted or b) objected to by the Examiner.
 Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).
 Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).
 11) The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.

Priority under 35 U.S.C. § 119

12) Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).
 a) All b) Some * c) None of:
 1. Certified copies of the priority documents have been received.
 2. Certified copies of the priority documents have been received in Application No. _____.
 3. Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).

* See the attached detailed Office action for a list of the certified copies not received.

Attachment(s)

1) Notice of References Cited (PTO-892) *2 sheets*
 2) Notice of Draftsperson's Patent Drawing Review (PTO-948)
 3) Information Disclosure Statement(s) (PTO-1449 or PTO/SB/08)
 Paper No(s)/Mail Date 10 sheets, 10/01/01, 9/7/01, 3/11/02

4) Interview Summary (PTO-413)
 Paper No(s)/Mail Date. _____.
 5) Notice of Informal Patent Application (PTO-152)
 6) Other: _____.

DETAILED ACTION

1. Applicant's election with traverse of Group I, claims 1-10 and species SEQ ID NO: 2, 5, 6, 7, 12-17, in the reply filed on June 10, 2004 is acknowledged. The traversal is on the ground(s) that for a restriction requirement to be proper, the Examiner must show (1) that the inventions defined by the restricted groups of claims are independent and distinct, and (2) that there would be a serious burden on the Examiner if restriction was not required. M.P.E.P. 803. Applicants assert that the Examiner has focused on only the first part of this two-part test. In order to properly restrict the groups, the Examiner needs to show that there would be a serious burden in examining the claims together. Applicants submit that no such serious burden exists, and respectfully submit that withdrawal of the restriction requirement as to the claims of Groups I, II and V is appropriate. Specifically, because the claims of each group are directed to compositions comprising at least one oligonucleotide comprising both an RNA region and a DNA region, wherein at least one terminus of the oligonucleotide comprises RNA, this subject matter must be searched for each group. Thus, a thorough search and examination of Groups I, II, and V together does not represent an undue burden and Applicants respectfully request that the Examiner reconsider this restriction requirement. See M.P.E.P. 803 ("If the search and examination of the entire application can be made without serious burden, the examiner must examine it on the merits, even though it includes claims to independent or distinct inventions." (emphasis added)).

Applicants expressly reserve their right, under M.P.E.P. 821.04, to add method claims to this application that depend from, or otherwise incorporate all limitations

of, the product claims of Groups I, II, and V for rejoinder with allowed product claims in this application.

This is not found persuasive. With regard to Applicants' assertions that there is no serious search burden to the Examiner to examine Groups I, II and V together, it is noted that a serious search burden is created because the Groups II and V would require additional search for the composition that comprises not only the RNA and DNA as set forth in the invention of Group I, but also the other ingredients claimed in Groups II and V, for example the antigen. The restriction Groups have acquired a separate status in the art as a separate subject for inventive effect and require independent searches. The search for each of the above inventions is not co-extensive particularly with regard to the literature search. A reference, which would anticipate the invention of one group would not necessarily anticipate or make obvious any of the other groups. Moreover, as to the question of burden of search, classification of subject matter is merely one indication of the burdensome nature of the search involved. The literature search, particularly relevant in this art, is not co-extensive and is much more important in evaluating the burden of search. Burden in examining materially different groups having materially different issues also exist. The restriction requirement between Groups I, II and V is maintained.

It is also noted that Applicants expressly reserve their right, under M.P.E.P. 821.04, to add method claims to this application that depend from, or otherwise incorporate all limitations of, the product claims of Group I for rejoinder with allowed product claims in this application. **Process claims that depend from or otherwise include all the limitations of the patentable product** will be entered as a matter of right if the amendment is presented prior to final rejection or allowance, whichever is earlier. Amendments, submitted after final rejection, are

governed by 37 CFR 1.116; amendments submitted after allowance are governed by 37 CFR 1.312.

In the event of rejoinder, the requirement for restriction between the product claims and the rejoined process claims will be withdrawn, and the rejoined process claims will be fully examined for patentability in accordance with 37 CFR 1.104. Thus, to be allowable, the rejoined claims must meet all criteria for patentability including the requirements of 35 U.S.C. 101, 102, 103, and 112. Until an elected product claim is found allowable, an otherwise proper restriction requirement between product claims and process claims may be maintained. Withdrawn process claims that are not commensurate in scope with an allowed product claim will not be rejoined. See “Guidance on Treatment of Product and Process Claims in light of *In re Ochiai*, *In re Brouwer* and 35 U.S.C. § 103(b),” 1184 O.G. 86 (March 26, 1996). Additionally, in order to retain the right to rejoinder in accordance with the above policy, Applicant is advised that the process claims should be amended during prosecution either to maintain dependency on the product claims or to otherwise include the limitations of the product claims. **Failure to do so may result in a loss of the right to rejoinder.**

Further, note that the prohibition against double patenting rejections of 35 U.S.C. 121 does not apply where the restriction requirement is withdrawn by the examiner before the patent issues. See MPEP § 804.01.

The requirement is still deemed proper and is therefore made FINAL.

2. Claims 11-17 have been withdrawn from further consideration pursuant to 37 CFR 1.142(b), as being drawn to a nonelected inventions, there being no

allowable generic or linking claim. Applicant timely traversed the restriction (election) requirement in the reply filed on June 10, 2004.

3. It is noted that elected SEQ ID NO: 2, 5, 6, and 7 have an effective filing date of June 7, 2000. These sequences were first disclosed in the provisional application 60/209797, filed June 7, 2000. Elected SEQ ID NO: 12-17 have an effective filing date of June 7, 2001. These sequences were first disclosed in the non-provisional application 09/874991, filed June 7, 2001.

4. Claims 8-10 are rejected under 35 U.S.C. 112, first paragraph, because the specification, while being enabling for *in vitro* use of the immunostimulatory compositions, does not reasonably provide enablement for *in vivo* use of the immunostimulatory compositions. The specification does not enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to use the invention commensurate in scope with these claims.

The claims are directed to an immunostimulatory composition comprising at least a first oligonucleotide and a second oligonucleotide, wherein both the first and second oligonucleotides each contain at least one RNA region and at least one DNA region, wherein at least one terminus of each oligonucleotide comprises RNA. Claims are also recite that each of the oligonucleotides of the immunostimulatory composition elicits a different immune stimulation profile. Claims are directed to an adjuvant comprising at least one oligonucleotide comprising both an RNA region and a DNA region, wherein at least one terminus of the oligonucleotide comprises RNA.

It is noted that the compositions contemplate both *in vitro* and *in vivo* use of these compositions.

The specification discloses *in vitro* methods (simulation of Th1 And Th2 type cytokine production, release of IL-6, release of IFN-gamma, B cell proliferation, stimulation of IgM secretion) using human peripheral lymphocytes and SEQ ID NO: 2, 5, 6, 7, HDR (hybrid DNA and RNA) and SEQ ID NO: 1 (only DNA, as a control) (see specification examples 2-7 on pages 34-47 and Tables 1-9). The specification teaches that the HDR does indeed stimulate different immune stimulation profiles, but only via *in vitro* means. Applicants speculate in the specification that, “it is expected that the HDRs of the invention, including mixtures of HDRs that elicit complementary patterns of activation, will provide correspondingly superior improvement to Th1 and Th2 responses in a patient as compared to DNA-based oligonucleotides.” (p. 36). However, the specification does not set forth any enablement for HDRs eliciting a different immune stimulation profile in an animal or patient or adjuvant activity in an animal or patient. Examples 8-10 (see pages 48-50 of the specification) set forth a prophetic protocol of how one of skill in the art would do these *in vivo* experiments, not that there were actually performed or that the immune stimulation was achieved in the same manner as for the *in vitro* experiments in Examples 2-7. Example 8 states, that “HDR injected mice will show increased levels of total IgM as opposed to the PBS injected controls.” (p. 48). Example 9 only shows *in vivo* use of SEQ ID NO: 2. Example 10 states that the “HDR injected animals *will show* elevated levels of anti-BSA ...”. (p. 50). The specification further asserts that the list of HDRs (see pages 51-64) are illustrative sequences have been selected in light of ODN sequences known in the art to posses immunostimulatory activity (innate, global,

cellular and/or humoral), and in light of the surprising observation reported herein that hybrid RNA-DNA ONDs (HDRs) possess robust immunostimulatory activity both *in vitro* and *in vivo*. Using the teachings of Examples 1-10, or other assays commonly used in the art, the skilled artisan will recognize that such HDRs, and all other HDR sequences within the scope of the invention can be assayed *in vitro* or *in vivo* for immunostimulatory activity (see p. 51). However, none of the sequences listed on pages 51-64 have been tested and they are not of the same structure as the tested SEQ ID NO: 2. SEQ ID NO: 2 is a RDR (meaning it has RNA at both termini and a DNA, CpG, center) while SEQ ID NO: 12-17 are DR (meaning only one terminus is RNA). It is not clear that a DR will function in the same manner as a RDR in either an *in vitro* or *in vivo* situation.

The state of the art is not clear with regard to the *in vivo* use of a hybrid RNA/DNA. Cong et al 2003 (BBRC, 2003, 310:1133-1139) teach that hybrid DNA/RNA provide similar *in vitro* results as set forth in the specification, but does not teach that this is yet possible *in vivo*. Cong et al indicate that these compounds *may* permit the development of therapeutic agents for use against cancer, asthma, allergy and infectious diseases and as adjuvant, but this has not yet been accomplished (pp.1137-1138). The art only teaches the *in vitro* methods of these hybrid compounds. In view of the lack of teaching and guidance in the specification and the unpredictability in the art with regard to *in vivo* use of these hybrids to elicit a different immune stimulation profile there would be undue experimentation necessary for a skilled artisan to practice the scope of the claimed invention.

5. The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless –

(b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

(e) the invention was described in (1) an application for patent, published under section 122(b), by another filed in the United States before the invention by the applicant for patent or (2) a patent granted on an application for patent by another filed in the United States before the invention by the applicant for patent, except that an international application filed under the treaty defined in section 351(a) shall have the effects for purposes of this subsection of an application filed in the United States only if the international application designated the United States and was published under Article 21(2) of such treaty in the English language.

6. Claims 1-10 are rejected under 35 U.S.C. 102(e) as being anticipated by Raz et al (6534062 6613751 or 6552006).

Raz et al, 6534062 for example, discloses a RNA/DNA hybrid and the DNA has a CpG motif (col. 7, l. 1-33; cols. 20-21). Raz et al discloses modifications of the phosphate backbone, which include a phosphorathioate modification (col. 7). The prior art anticipates the claimed invention.

With regard to claims 9, “that each of the oligonucleotides of the immunostimulatory composition elicits a different immune stimulation profile” and claim 10, “an adjuvant comprising at least one oligonucleotide comprising both an RNA region and a DNA region, wherein at least one terminus of the oligonucleotide comprises RNA”, it is noted that the prior art discloses the structural components of the claimed invention. The properties defined in claim 9 are believed to inherent. With regard to claim 10, the recitation of “adjuvant” is viewed as intended use. A recitation of the intended use of the claimed invention must result in a structural difference between the claimed invention and the prior

art in order to patentably distinguish the claimed invention from the prior art. If the prior art structure is capable of performing the intended use, then it meets the claim. In a claim drawn to a process of making, the intended use must result in a manipulative difference as compared to the prior art. See *In re Casey*, 152 USPQ 235 (CCPA 1967) and *In re Otto*, 136 USPQ 458, 459 (CCPA 1963).

Since the Office does not have the facilities for examining and comparing applicants' compositions with the compositions of the prior art, the burden is on applicant to show a novel or unobvious differences between the claimed product and the product of the prior art (i.e., that the compositions of the prior art does not possess the same material structural and functional characteristics of the claimed compositions) See In re Best, 562 F.2d 1252, 195 USPQ 430 (CCPA 1977) and In re Fitzgerald et al., 205 USPQ 594.

7. Claims 1-10 are rejected under 35 U.S.C. 102(b) as being anticipated by Fosnaugh et al 1989 (Molecular and Cellular Biology, Nov. 1989, 9/11:5215-5218).

Fosnaugh et al disclose the elected SEQ ID NO: 7. It is noted that the claims are directed to an immunostimulatory composition comprising: at least one oligonucleotide comprising both an RNA region and a DNA region, wherein at least one terminus of the oligonucleotide comprises RNA. The prior art discloses the RNA at either or both termini and a DNA sequence that has a CpG motif as claimed.

With regard to claims 9, "that each of the oligonucleotides of the immunostimulatory composition elicits a different immune stimulation profile" and claim 10, "an adjuvant comprising at least one oligonucleotide comprising

both an RNA region and a DNA region, wherein at least one terminus of the oligonucleotide comprises RNA”, it is noted that the prior art discloses the structural components of the claimed invention. The properties defined in claim 9 are believed to inherent. With regard to claim 10, the recitation of “adjuvant” is viewed as intended use. A recitation of the intended use of the claimed invention must result in a structural difference between the claimed invention and the prior art in order to patentably distinguish the claimed invention from the prior art. If the prior art structure is capable of performing the intended use, then it meets the claim. In a claim drawn to a process of making, the intended use must result in a manipulative difference as compared to the prior art. See *In re Casey*, 152 USPQ 235 (CCPA 1967) and *In re Otto*, 136 USPQ 458, 459 (CCPA 1963).

Since the Office does not have the facilities for examining and comparing applicants' compositions with the compositions of the prior art, the burden is on applicant to show a novel or unobvious differences between the claimed product and the product of the prior art (i.e., that the compositions of the prior art does not possess the same material structural and functional characteristics of the claimed compositions) See *In re Best*, 562 F.2d 1252, 195 USPQ 430 (CCPA 1977) and *In re Fitzgerald et al.*, 205 USPQ 594.

8. Claims 1-10 are rejected under 35 U.S.C. 102(e) as being anticipated by Lees et al 6632923 or Dumas et al 6639063.

Lees et al discloses the elected species SEQ ID NO: 16 (see SEQ ID NO: 51 of the patent).

Dumas et al discloses the elected species SEQ ID NO: 7 (see SEQ ID NO: 11708 of the patent).

With regard to claims 9, “that each of the oligonucleotides of the immunostimulatory composition elicits a different immune stimulation profile” and claim 10, “an adjuvant comprising at least one oligonucleotide comprising both an RNA region and a DNA region, wherein at least one terminus of the oligonucleotide comprises RNA”, it is noted that the prior art discloses the structural components of the claimed invention. The properties defined in claim 9 are believed to inherent. With regard to claim 10, the recitation of “adjuvant” is viewed as intended use. A recitation of the intended use of the claimed invention must result in a structural difference between the claimed invention and the prior art in order to patentably distinguish the claimed invention from the prior art. If the prior art structure is capable of performing the intended use, then it meets the claim. In a claim drawn to a process of making, the intended use must result in a manipulative difference as compared to the prior art. See *In re Casey*, 152 USPQ 235 (CCPA 1967) and *In re Otto*, 136 USPQ 458, 459 (CCPA 1963).

Since the Office does not have the facilities for examining and comparing applicants' compositions with the compositions of the prior art, the burden is on applicant to show a novel or unobvious differences between the claimed product and the product of the prior art (i.e., that the compositions of the prior art does not possess the same material structural and functional characteristics of the claimed compositions) See *In re Best*, 562 F.2d 1252, 195 USPQ 430 (CCPA 1977) and *In re Fitzgerald et al.*, 205 USPQ 594.

9. Claims 1-10 are rejected under 35 U.S.C. 102(b) as being anticipated by Accession No. AP000792 and AP000728 published in the Database 1999.

Accession No. AP000792 and AP000728 disclose elected species SEQ ID NO: 6 (see the attached sequence search printout).

With regard to claims 9, “that each of the oligonucleotides of the immunostimulatory composition elicits a different immune stimulation profile” and claim 10, “an adjuvant comprising at least one oligonucleotide comprising both an RNA region and a DNA region, wherein at least one terminus of the oligonucleotide comprises RNA”, it is noted that the prior art discloses the structural components of the claimed invention. The properties defined in claim 9 are believed to inherent. With regard to claim 10, the recitation of “adjuvant” is viewed as intended use. A recitation of the intended use of the claimed invention must result in a structural difference between the claimed invention and the prior art in order to patentably distinguish the claimed invention from the prior art. If the prior art structure is capable of performing the intended use, then it meets the claim. In a claim drawn to a process of making, the intended use must result in a manipulative difference as compared to the prior art. See *In re Casey*, 152 USPQ 235 (CCPA 1967) and *In re Otto*, 136 USPQ 458, 459 (CCPA 1963).

Since the Office does not have the facilities for examining and comparing applicants' compositions with the compositions of the prior art, the burden is on applicant to show a novel or unobvious differences between the claimed product and the product of the prior art (i.e., that the compositions of the prior art does not possess the same material structural and functional characteristics of the claimed compositions) See *In re Best*, 562 F.2d 1252, 195 USPQ 430 (CCPA 1977) and *In re Fitzgerald et al.*, 205 USPQ 594.

10. Claims 1-10 are rejected under 35 U.S.C. 102(b) as being anticipated by Bonaldo et al Genome Research, 1996, 6/9:791-806.

Bonaldo et al discloses the elected species SEQ ID NO: 6 (see the attached sequence search printout).

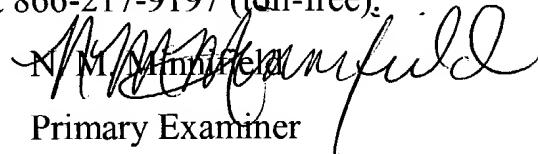
With regard to claims 9, "that each of the oligonucleotides of the immunostimulatory composition elicits a different immune stimulation profile" and claim 10, "an adjuvant comprising at least one oligonucleotide comprising both an RNA region and a DNA region, wherein at least one terminus of the oligonucleotide comprises RNA", it is noted that the prior art discloses the structural components of the claimed invention. The properties defined in claim 9 are believed to inherent. With regard to claim 10, the recitation of "adjuvant" is viewed as intended use. A recitation of the intended use of the claimed invention must result in a structural difference between the claimed invention and the prior art in order to patentably distinguish the claimed invention from the prior art. If the prior art structure is capable of performing the intended use, then it meets the claim. In a claim drawn to a process of making, the intended use must result in a manipulative difference as compared to the prior art. See *In re Casey*, 152 USPQ 235 (CCPA 1967) and *In re Otto*, 136 USPQ 458, 459 (CCPA 1963).

Since the Office does not have the facilities for examining and comparing applicants' compositions with the compositions of the prior art, the burden is on applicant to show a novel or unobvious differences between the claimed product and the product of the prior art (i.e., that the compositions of the prior art does not possess the same material structural and functional characteristics of the claimed compositions) See *In re Best*, 562 F.2d 1252, 195 USPQ 430 (CCPA 1977) and *In re Fitzgerald et al.*, 205 USPQ 594.

11. No claims are allowed.
12. The prior art made of record and not relied upon is considered pertinent to applicant's disclosure.
13. Any inquiry concerning this communication or earlier communications from the examiner should be directed to N. M. Minnifield whose telephone number is 571-272-0860. The examiner can normally be reached on M-F (8:00-5:30) Second Friday Off.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Lynette R.F. Smith can be reached on 571-272-0864. The fax phone number for the organization where this application or proceeding is assigned is 703-872-9306.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see <http://pair-direct.uspto.gov>. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free).


N. M. Minnifield
Primary Examiner

Art Unit 1645

NMM

August 31, 2004

FILE REFERENCE: GENSET.054PR2
 CURRENT APPLICATION NUMBER: US/09/621,976
 CURRENT FILING DATE: 2000-07-21
 NUMBER OF SEQ ID NOS: 19335
 SOFTWARE: Patent.pn
 SEQ ID NO: 12427
 LENGTH: 129
 ORGANISM: Homo sapiens
 US-09-621-976-12427

Query Match 97.3%; Score 29.2; DB 4; Length 129;
 Best Local Similarity 93.3%; Pred. No. 13; DB 4; Length 254;
 Matches 28; Conservative 2; Mismatches 0; Indels 0; Gaps 0;
 SEQ ID NO: 12427

Query 1 AAAAAAAAAAAACGAAAGAAAAAA 30
 Db 97 AAAAAAAAAAAACGAAAGAAAAAA 126

RESULT 5
 US-09-621-976-14176
 Sequence 14176, Application US/09621976
 ; GENERAL INFORMATION:
 ; Patent No. 6639063
 ; APPLICANT: Dumas Milne Edwards, J.B.
 ; APPLICANT: Jobert, S.
 ; APPLICANT: Giordano, J.Y.
 ; TITLE OF INVENTION: ESTs and Encoded Human Proteins.
 ; FILE REFERENCE: GENSET.054PR2
 ; CURRENT APPLICATION NUMBER: US/09/621,976
 ; CURRENT FILING DATE: 2000-07-21
 ; NUMBER OF SEQ ID NOS: 19335
 ; SOFTWARE: Patent.pn
 ; SEQ ID NO: 14176

Query Match 94.7%; Score 28.4; DB 4; Length 77;
 Best Local Similarity 96.7%; Pred. No. 20; DB 4; Length 77;
 Matches 29; Conservative 0; Mismatches 1; Indels 0; Gaps 0;
 SEQ ID NO: 14176

Query 1 AAAAAAAAAAAACGAAAGAAAAAA 30
 Db 47 AAAAAAAAAAAACGAAAGAAAAAA 76

RESULT 6
 US-09-621-976-13606
 Sequence 13606, Application US/09621976
 ; GENERAL INFORMATION:
 ; Patent No. 6639063
 ; APPLICANT: Dumas Milne Edwards, J.B.
 ; APPLICANT: Jobert, S.
 ; APPLICANT: Giordano, J.Y.
 ; TITLE OF INVENTION: ESTs and Encoded Human Proteins.
 ; FILE REFERENCE: GENSET.054PR2
 ; CURRENT APPLICATION NUMBER: US/09/621,976
 ; CURRENT FILING DATE: 2000-07-21
 ; NUMBER OF SEQ ID NOS: 19335
 ; SOFTWARE: Patent.pn
 ; SEQ ID NO: 13606
 ; LENGTH: 183
 ; TYPE: DNA
 ; ORGANISM: Homo sapiens
 ; US-09-621-976-13606

Query Match 94.7%; Score 28.4; DB 4; Length 163;
 Best Local Similarity 96.7%; Pred. No. 19; DB 4; Length 163;
 Matches 28; Conservative 0; Mismatches 1; Indels 0; Gaps 0;
 SEQ ID NO: 13606

Query 1 AAAAAAAAAAAACGAAAGAAAAAA 30
 Db 132 AAAAAAAAAAAACGAAAGAAAAAA 161

RESULT 7
 US-09-621-976-10240
 Sequence 10240, Application US/09621976
 ; GENERAL INFORMATION:
 ; Patent No. 6639063
 ; APPLICANT: Dumas Milne Edwards, J.B.
 ; APPLICANT: Jobert, S.
 ; APPLICANT: Giordano, J.Y.
 ; TITLE OF INVENTION: ESTs and Encoded Human Proteins.
 ; FILE REFERENCE: GENSET.054PR2
 ; CURRENT APPLICATION NUMBER: US/09/621,976
 ; CURRENT FILING DATE: 2000-07-21
 ; NUMBER OF SEQ ID NOS: 19335
 ; SOFTWARE: Patent.pn
 ; SEQ ID NO: 17033
 ; LENGTH: 254
 ; TYPE: DNA
 ; ORGANISM: Homo sapiens
 ; US-09-621-976-17033

FEATURES	Source	Location/Qualifiers
1.	.30	/organism="synthetic construct"
/mol_type="unassigned DNA"		
/db_xref="taxon:32630"		
/note="Description of Combined DNA/RNA Molecule: Synthetic DNA/RNA Oligonucleotide-Synthetic DNA/RNA oligonucleotide"		
ORIGIN		
Query Match	Score 30; DB 6; Length 30;	
Best Local Similarity	100.0% ; Pred. No. 11;	
Matches 30; Conservative 0; Mismatches 0; Indels 0; Gaps 0;		
Qy	1 AAAAAAAAAACCCGGAAAGAAAAAA 30	
Db	1 AAAAAAAAAACCCGGAAAGAAAAAA 30	
RESULT 2		
AP000792	AP000792 DNA linear HTG 30-MAY-2000	
DEFINITION	Homo sapiens chromosome 11 clone RP11-792M23 map 11q14, WORKING DRAFT	
ACCESSION	AP000792	
VERSION	AP000792.2	
KEYWORDS	HTGS; PHASE1; HTGS DRAFT.	
ORGANISM	Homo sapiens (human)	
REFERENCE		
AUTHORS	Hattori, M., Ishii, K., Toyoda, A., Taylor, T.D., Hong-Seeq, P., Fujiyama, A., Yada, T., Totoki, Y., Watanabe, H. and Sakaki, Y.	
TITLE	Homo sapiens 157,712 genomic DNA of 11q14	
JOURNAL	Published Only in DataBase (1999)	
REFERENCE		
AUTHORS	Hattori, M., Ishii, K., Toyoda, A., Taylor, T.D., Hong-Seeq, P., Fujiyama, A., Yada, T., Totoki, Y., Watanabe, H. and Sakaki, Y.	
TITLE	Direct Submission	
JOURNAL	Submitted (29-NOV-1999) Masahira Hattori, The Institute of Physical and Chemical Research (RIKEN), Genomic Sciences Center (GSC); Kitasato Univ., 1-15-1 Kitasato, Sagamihara, Kanagawa 228-8555, Japan (E-mail:hattori@gsc.riken.go.jp, URL: http://hgp.ssc.riken.go.jp/, Fax: 81-42-778-9224)	
COMMENT	On May 31, 2000 this sequence version replaced gi:6997629.	
Center: RIKEN Genomic Sciences Center (GSC)		
Center code: RIKEN		
Web site: http://hgp.ssc.riken.go.jp/		
Contact: hattori@gsc.riken.go.jp		
----- Project Information		
Center Project name: HumDwarf1		
Center Clone name: RP11-792M23		
----- Summary Statistics		
Sequencing vector: PCR Products, 100% of reads		
Chemistry: Dye-terminator Br-amersham; 100% of reads		
Assembly program: Phrap; version .990329		
Consensus quality: 143304 bases at least Q40		
Consensus quality: 153360 bases at least Q30		
Consensus quality: 156339 bases at least Q20		
Insert size: 157812; sum-of-contigs		
Quality coverage: 4.7x in Q20 bases; sum-of-contigs		

NOTE: This is a 'working draft' sequence. It currently consists of 20 contigs. The true order of the pieces is not known and their order in this sequence record is arbitrary. Gaps between the contigs are represented as runs N, but the exact sizes of the gaps are unknown. This record will be updated with the finished sequence as soon as it is available and the accession number will be preserved.		
1	20543 contig of 20543 bp in length	
2	41423 contig of 20780 bp in length	

FEATURES
Source

/organism="Homo sapiens"
/mol_type="genomic DNA"
/db_xref="taxon:9606"
/chitosome="11"
/map="11q14"

Sequence undated (26-May-2000)

* NOTE: This is a 'working draft' sequence. It currently consists of 20 contigs. The true order of the pieces is not known and their order in this sequence record is arbitrary. Gaps between the contigs are represented as runs of N, but the exact sizes of the gaps are unknown. This record will be updated with the finished sequence as soon as it is available and the accession number will be preserved.

* 1 20543: contig of 20543 bp in length

* 2 20643: gap of 100 bp

* 3 41423: contig of 20780 bp in length

* 4 41424: gap of 100 bp

* 5 41523: gap of 100 bp

* 6 41524: contig of 16549 bp in length

* 7 58072: gap of 100 bp

* 8 58073: gap of 100 bp

* 9 58173: gap of 100 bp

* 10 74856: contig of 16684 bp in length

* 11 74956: gap of 100 bp

* 12 74957: contig of 15811 bp in length

* 13 90767: gap of 100 bp

* 14 90868: contig of 12673 bp in length

* 15 10354: gap of 100 bp

* 16 10364: gap of 100 bp

* 17 112882: contig of 9242 bp in length

* 18 112883: gap of 100 bp

* 19 112884: contig of 5500 bp in length

* 20 118483: gap of 100 bp

* 21 118484: gap of 100 bp

* 22 118485: contig of 7596 bp in length

* 23 126179: gap of 100 bp

* 24 132359: contig of 6081 bp in length

* 25 132360: gap of 100 bp

* 26 132459: contig of 198 bp in length

* 27 137425: contig of 4966 bp in length

* 28 137525: gap of 100 bp

* 29 141745: contig of 4220 bp in length

* 30 141845: gap of 100 bp

* 31 145942: contig of 4097 bp in length

* 32 145943: gap of 100 bp

* 33 145042: gap of 100 bp

* 34 146143: contig of 2198 bp in length

* 35 148440: gap of 100 bp

* 36 148340: gap of 100 bp

* 37 148341: gap of 100 bp

* 38 143341: contig of 2334 bp in length

* 39 150675: gap of 100 bp

* 40 150775: gap of 1560 bp in length

* 41 152334: gap of 100 bp

* 42 152435: contig of 2037 bp in length

* 43 154471: gap of 100 bp

* 44 154572: gap of 2086 bp in length

* 45 155657: gap of 100 bp

* 46 156758: contig of 1791 bp in length

* 47 156759: gap of 100 bp

* 48 158548: gap of 100 bp

* 49 158649: contig of 1064 bp in length

1. 159712: Location/Qualifiers

```

RESULT 3
P000728          AP000728          150594 bp   DNA  linear  HTG 30-MAY-2000
          OCTU          Homo sapiens chromosome 11 clone RP11-699G11 map 11q14, WORKING
          EFPIINTCN

```

Sequence updated (26-May-2000).

* NOR: This is a 'working draft' sequence. It currently consists of 48 contigs. The true order of the pieces is not known and their order in this sequence record is arbitrary. Gaps between the contigs are represented as runs of N, but the exact sizes of the gaps are unknown. This record will be updated with the finished sequence as soon as it is available and the accession number will be preserved.

1 7885: contig of 7885 bp in length

2 7985: gap of 100 bp

3 7986: 1631: contig of 8325 bp in length

4 1631: 1640: gap of 100 bp

5 1641: 22081: contig of 5678 bp in length

6 22089: 22188: gap of 100 bp

7 22189: 28354: contig of 6170 bp in length

8 28359: 28448: gap of 100 bp

9 28449: 34160: contig of 5702 bp in length

10 34265: 39101: gap of 100 bp

11 39101: 39201: contig of 4841 bp in length

12 39201: 39202: gap of 100 bp

13 44288: 44388: contig of 5087 bp in length

14 44289: 44389: gap of 100 bp

15 50127: 50227: contig of 5739 bp in length

16 50228: 55556: gap of 100 bp

17 55556: 55656: contig of 5329 bp in length

18 55657: 58778: contig of 3122 bp in length

19 58879: 58878: gap of 100 bp

20 58879: 62612: contig of 3734 bp in length

21 62613: 62713: gap of 100 bp

22 62713: 66185: contig of 3472 bp in length

23 66285: 70337: gap of 100 bp

24 70338: 70838: contig of 4453 bp in length

25 75173: 75273: gap of 100 bp

26 75273: 79442: contig of 4335 bp in length

27 79442: 83749: gap of 100 bp

28 83749: 83580: contig of 3837 bp in length

29 83580: 86337: gap of 100 bp

30 86337: 86446: contig of 2657 bp in length

31 86446: 89885: gap of 100 bp

32 89885: 93305: contig of 3449 bp in length

33 93305: 93406: gap of 100 bp

34 93406: 96390: contig of 3320 bp in length

35 96390: 99799: gap of 100 bp

36 99799: 105420: contig of 3210 bp in length

37 105420: 105520: gap of 100 bp

38 105520: 108049: contig of 3253 bp in length

39 108049: 108149: gap of 100 bp

40 108149: 102153: contig of 320 bp in length

41 102153: 105419: contig of 3167 bp in length

42 105419: 110752: gap of 100 bp

43 110752: 114852: contig of 3512 bp in length

44 114852: 114364: gap of 100 bp

45 114364: 115521: contig of 1058 bp in length

46 115521: 115621: gap of 100 bp

47 115621: 117656: contig of 2035 bp in length

48 117656: 119830: gap of 100 bp

49 119830: 149244: contig of 2074 bp in length

50 149244: 143058: gap of 100 bp

51 143058: 144610: contig of 1452 bp in length

52 144610: 145753: gap of 100 bp

53 145753: 145891: contig of 1694 bp in length

54 145891: 146912: gap of 100 bp

55 146912: 147012: contig of 1446 bp in length

56 147012: 148183: gap of 100 bp

57 148183: 149244: contig of 1759 bp in length

58 149244: 150594: gap of 100 bp

59 150594: 152189: contig of 2053 bp in length

60 152189: 153536: gap of 100 bp

61 153536: 153637: contig of 1832 bp in length

62 153637: 153937: gap of 100 bp

63 153937: 154592: contig of 100 bp

64 154592: 155396: gap of 100 bp

65 155396: 157081: contig of 1592 bp in length

66 157081: 157187: gap of 100 bp

67 157187: 158305: contig of 1118 bp in length

68 158305: 158405: gap of 100 bp

69 158405: 159438: contig of 1033 bp in length

70 159438: 159538: gap of 100 bp

71 159538: 159539: contig of 2112 bp in length

72 159539: 161650: gap of 100 bp

73 161650: 164750: contig of 1308 bp in length

74 164750: 164751: gap of 100 bp

75 164751: 164753: contig of 1452 bp in length

76 164753: 164754: gap of 100 bp

77 164754: 164755: contig of 1053 bp in length

78 164755: 164756: gap of 100 bp

79 164756: 164757: contig of 1048 bp in length

80 164757: 164758: gap of 100 bp

81 164758: 164759: contig of 1172 bp in length

82 164759: 164823: gap of 100 bp

83 164823: 164824: contig of 1172 bp in length

84 164824: 164825: gap of 100 bp

85 164825: 164826: contig of 1172 bp in length

86 164826: 164827: gap of 100 bp

87 164827: 164828: contig of 1172 bp in length

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91 164831: 164832: contig of 1172 bp in length

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223 164963: 164964: contig of 1172 bp in length

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260 165000: 165001: gap of 100 bp

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262 165002: 165003: gap of 100 bp

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272 165012: 165013: gap of 100 bp

273 165013: 165014: contig of 1172 bp in length

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275 165015: 165016: contig of 1172 bp in length

276 165016: 165017: gap of 100 bp

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283 165023: 165024: contig of 1172 bp in length

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322 165062: 165063: gap of 100 bp

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339 165079: 165080: contig of 1172 bp in length

340 165080: 165081: gap of 100 bp

341 165081: 165082: contig of 1172 bp in length

342 165082: 165083: gap of 100 bp

343 165083: 16508

Please mail w/ Action

COMMENT	Fax:81-75-705-1113 Ciona intestinalis cDNA Project (URL: http://ghoz.volc.kyoto-u.ac.jp/index1.html).	
FEATURES		
ORIGIN		
RESULT		
COMMENT	1. •1458 /organism="Ciona intestinalis" /mol_type="mRNA" /db_xref="taxon:719" /clone="cieg056p11"	
FEATURES	Query Match Best Local Similarity 100.0%; P-Value 0.3 6e+02; Matches 30; Conservative 0; Mismatches 0; Ind	
ORIGIN	Qy 1 AAAAAAAAAAAGCAAAAGAAAAAA 30 Dd 1425 AAAAAAAGAAAGCAAAAGAAAAAA 1454	
RESULT	6	
LOCUS	AY080787	1538 bp mRNA line
DEFINITION	Arabidopsis thaliana putative enoyl-CoA hydratase	
ACCESSION	AY080787	complete cds.
VERSION	AY080787.1	
KEYWORDS	GI:19424018	
SOURCE	PLI cDNA	
ORGANISM	Arabidopsis thaliana (thale cress)	
REFERENCE	1	
AUTHORS	Yamada,K., Liu,S.X., Sakano,H., Pham,P.K., Bans	
VERSION	Goldsmith,A.D., Lee,J.M., Quach,H.L., Toriumi,	
KEYWORDS	Carninci,P., Chen,H., Cheuk,R., Hayashizaki,Y.	
SOURCE	Jones,T., Kamiya,A., Karin-Neumann,G., Kawai,	
ORGANISM	Lin,J., Miranda,M., Naruse,M., Nguyen,M., Pa	
REFERENCE	Satou,M., Seki,M., Shinn,P., Southwick,A., Shi	
AUTHORS	Davis,R.W., Ecker,J.R., and Theologis,A.	
VERSION	Arabidopsis Full Length cDNA Clones	
KEYWORDS	Unpublished	
SOURCE	2 (bases 1 to 1538)	
ORGANISM	Yamada,K., Banh,J., Chan,M.M., Chang,C.H., Cha	
REFERENCE	Deng,J.M., Goldsmith,A.D., Lee,J.M., Onodera,C	
AUTHORS	Tang,C.C., Toriumi,M., Wu,H.C., Yamamoto,Y.	
VERSION	Carninci,P., Chen,H., Cheuk,R., Hayashizaki,Y.	
KEYWORDS	Jones,T., Kamiya,A., Karin-Neumann,G., Kawai,	
SOURCE	Lin,J., Meyers,M.C., Miranda,M., Naruse,M.,	
ORGANISM	Sakurai,T., Satou,M., Seki,M., Shinn,P., South	
REFERENCE	Shinozaki,K., Davis,R.W., Ecker,J.R., and Theol	
AUTHORS	Submitted (19-FEB-2002) Plant Gene Expression	
VERSION	Street, Albany, CA 94710, USA	
KEYWORDS	RIKEN Genomic Sciences Center (GSC) members ca	
SOURCE	collection and clustering of R AFLP cDNAs (RAFL	
ORGANISM	Arabidopsis Full Length cDNA.)	
REFERENCE	Satou,M., Kamiya,A., Sakurai,T., Carninci,P.,	
AUTHORS	Hayashizaki,Y., and Shinozaki,K.	
VERSION	The Salk, Stanford, PGEC (SSP) Consortium memb	
KEYWORDS	sequencing and annotation of the R AFLP cDNAs: Y	
SOURCE	Chan,M.M., Chang,C.H., Chang,E., Dale,J.M., De	
ORGANISM	Goldsmith,A.D., Lee,J.M., Onodera,C.S., Quach,	
REFERENCE	Toriumi,M., Wu,H.C., Yamamoto,Y., Yu,G., Bowes	
AUTHORS	Cheuk,R., Jones,T., Karin-Neumann,G., Kim,C.,	
VERSION	Meyers,M.C., Miranda,M., Nguyen,M., Palm,C.J.,	
KEYWORDS	Southwick,A., Davis,R.W., Ecker,J.R., and Theol	
SOURCE	Yamada,K., (SSP/PGEC) and Seki M. (RIKEN GSC) c	
ORGANISM	Yamada,K., Shinozaki,K., (RIKEN GSC) and Theol	
REFERENCE	contributed equally to this work as PIs.	

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Page

oligonucleotide that was used to prime the synthesis of first strand cDNA and therefore this may represent a bona fide poly A signal. The sequence tag present in the cDNA between the Not I site and the Oligo-dT track served to verify it as a clone from the hippocampus tissue cDNA Library Preparation: M.B. Soares Lab CLONE LIBRARY: Researchers may obtain BMAP cDNA clones from RESEARCH GENETICS. It should be noted that Bento Soares is generating a small number of additional special non-redundant arrays of cDNAs whose availability will be considered under appropriate and limited collaborative arrangements. The following repetitive elements were found in this cDNA sequence: 1-5, >POLY A#Simple_repeat 143-277, >POLY A#Simple_repeat 361-432, >GC_rich#Low complexity seq primer: M13 Forward POLYA=Yes Location/Qualifiers 1. .467 /mol_type="mRNA" /strain="C57BL/6J" /db_xref="taxon:10090" /clone="T1-M-B21-bec-d-04-0-UI" /dev_stage="27-32 days" /lab_host="DH10B (Life Technologies)" /clone_lib="NIH BMAP MH12 S1" /note="Vector: pT7T3D-Pac (Pharmacia) with a modified polylinker; Site 1: Not I; Site 2: Eco RI; The NIH BMAP MH12 S1 library is a subtracted library derived from NIH BMAP MH12. NIH BMAP MH12 is a library derived from mouse hippocampus tissue. For a detailed description of the library from which this clone was derived, please visit our web site at brainest.eng.uiowa.edu. TAG TISSUE=hippocampus TAG LIB-NIH BMAP MH12 S1 TAG SOURCE=NIH BMAP MH12 S1

ATURES
Source

QIGIN
 Query Match 100.0%; Score 30; DB 10; Length 467;
 Best Local Similarity 100.0%; Pred. No. 6.1e+03;
 Matches 30; Conservative 0; Mismatches 0; Indels 0; Gaps 0
 1 AAAAAAAAAACCCGGAAAAAAAAAAA 30
 207 AAAAAAAAAACCCGGAAAAAAAAAA 236
 RSLT 2
 BU944962 887 bp mRNA linear EST 18-OCT-2001
 AGENCOURT 10546565 NIH_MGC_141 Homo sapiens cDNA clone
 IMAGE:6731765', mRNA sequence.
 BU944962
 BU944962.1 GI:24133781
 EST.
 Homo sapiens (human)
 Homo sapiens
 Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
 Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
 ACCESSION
 BU944962
 VERSION
 1
 FIELDS
 SOURCE
 ORGANISM
 FERENC
 AUTHORS
 TITLE
 JOURNAL
 COMMENT
 Tissue Procurement: NCI
 cDNA Library Preparation: Michael Brownstein Laboratory
 cDNA Library Arrayed by: The I.M.A.G.E. Consortium (LiNL)
 DNA Sequencing by: Agencourt Bioscience Corporation
 Clone distribution: MGC clone distribution information can be
 found through the I.M.A.G.E. Consortium/LiNL at:
 http://image.linnl.gov
 Plate: ILLCM3062 row: f column: 23

FEATURES	Location/Qualifiers
1. . 887	/organism="Homo sapiens"
	/mol_type="mRNA"
	/db_xref="taxon:9606"
	/clone lib="NIH MGC 141"
	/note="Vector: PNR-LIB; Site-1: SfiI (ggccattatggcc) ; Site-2: SfiI (ggccggcttggcc); Double-stranded cDNA was prepared from a pool of 40 cell line polyA+ RNAs (bladder - 2%, blood - 33.4%, brain - 5.6%, breast - 12.5%, colon - 4%, connective tissue - 1.4%, eye - 1%, intestine - 2.6%, kidney - 2.2%, liver - 5.7%, lung - 10.8%, NK-cell - 2.6%, ovary - 4%, pharynx - 2.5%, prostate - 4.3%, salivary gland - 1.3%, and skin - 2.3%). 5', and 3', adaptors were used in cloning as follows:
	5'-AAGCAGTGGATCAAGCGAGTCATGGCGATCATGGCGGG-3', and 5'-ATTCTPAGGCGCGGCGGCGGCGACATG(310)NN-3'. Full-length enriched library was constructed using the Clontech Creator SMART kit and size-selected to contain the 0.2-0.5 kb size fraction (other fractions present in NIH MGC 142). Library created in the laboratory of M. Brownstein (NIH, NIH). Note: this is a NIH_MGC Library."
ORIGIN	
	Query Match 100.0% ; Score 30; DB 13; Length 887;
	Best Local Similarity 100.0% ; Pred. No. 3.7e+03;
	Matches 30; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
Qy	1 AAAA.....AAACCCGGAAAAAAA 30
Db	341 AAAA.....AAACCCGGAAAAAAA 370
RESULT 3	
AL514409	582 bp mRNA linear EST 08-MAY-2003
LOCUS	AL514409 Homo sapiens NEUROBLASTOMA Homo sapiens cDNA clone CLOBB009ZC11 3-PRIME, mRNA sequence.
DEFINITION	
AL514409	AL514409
ACCESSION	AL514409.2 GL:30464294
VERSION	
KEYWORDS	
SOURCE	Homo sapiens (human)
ORGANISM	Homo sapiens
Bukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Primates; Catarrhini; Hominoidea; Homo	
REFERENCE	1 (bases 1 to 582)
AUTHORS	Li W.B., Gruber C., Jessee J. and Polayes D.
TITLE	Full-length cDNA libraries and normalization
JOURNAL	Unpublished (2001)
COMMENT	On Feb 13, 2001 this sequence version replaced gi:12777903.
FEATURES	Contact: Genoscope - Centre National de Sequentage
source	BP 191 91006 EVRY cedex - France
	Email: segret@genoscope.cns.fr Web: www.genoscope.cns.fr
	Library was constructed by Life Technologies, a division of Invitrogen. This sequence belongs to sequence cluster 10492.f. For more information about this cluster, see http://www.Genoscope.cns.fr/cgi-bin/cluster.cgi?seq=CL0BB009ZC11&cluster=10492.f . Contact: Feng Liang Email: filang@lifetech.com URL: http://fulllength.invitrogen.com/ InvitroGen Corporation 1600 Parady Avenue Genoscope sequence ID : CL0BB009ZC11fPI.
	Location/Qualifiers
1. . 582	/organism="Homo sapiens"
	/mol_type="mRNA"
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	/clone lib="CL0BB009ZC11"
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